

Amendments to the Claims

Claims 1-63. (Cancelled).

64. (Currently amended) A method of making one or more cDNA molecules, comprising:

- (a) mixing one or more RNA molecules with (i) one or more polypeptides having reverse transcriptase activity and (ii) at least one primer-adapter nucleic acid molecule, wherein the at least one primer-adapter nucleic acid molecule comprises one or more ligands and one or more cleavage sites, to form a mixture;
- (b) incubating the mixture under conditions sufficient to make one or more double stranded cDNA molecules, wherein one or more of the cDNA molecules comprise at least one primer-adapter nucleic acid molecule;
- (c) contacting one or more of the cDNA molecules with at least one hapten to produce one or more hapten-cDNA molecule complexes; and
- (d) cleaving one or more of the complexes with not more than one enzyme that cleaves one or more of the complexes at one or more cleavage sites within the primer-adapter, to produce one or more cleaved cDNA molecules; and
- (e) inserting or ligating one or more of the cleaved cDNA molecules into one or more vectors.

65. (Previously presented) The method according to claim 64, further comprising isolating one or more of the hapten-cDNA molecule complexes.

66. (Previously presented) The method according to claim 64, wherein at least one hapten is bound to a solid support.

67. (Previously presented) The method according to claim 66, wherein the solid support is selected from the group consisting of nitrocellulose, diazocellulose, glass, polystyrene, polyvinylchloride, polypropylene, polyethylene, dextran, Sepharose, agar, starch, nylon, beads and microtitre plates.

68. (Previously presented) The method according to claim 66, wherein the solid support is a magnetic bead.

69. (Cancelled).

70. (Currently amended) The method according to claim 69 64, wherein the restriction enzyme is *NotI*.

71. (Currently amended) The method according to claim 69 64, wherein one or more of the cleaved cDNA molecules comprises one sticky end and one blunt end.

72. (Previously presented) The method according to claim 71, wherein the sticky end is a *NotI* sticky end and the vector has a *NotI* compatible end and a blunt end.

73. (Previously presented) The method according to claim 64, wherein the one or more polypeptides are selected from the group consisting of a Moloney Leukemia Virus (M-MLV) reverse transcriptase, a Rous Sarcoma Virus (RSV) reverse transcriptase, an Avian Myeloblastosis Virus (AMV) reverse transcriptase, a Rous Associated Virus (RAV) reverse transcriptase, a Myeloblastosis Associated Virus (MAV) reverse transcriptase, a Human Immunodeficiency Virus (HIV) reverse transcriptase, a retroviral reverse transcriptase, a retrotransposon reverse transcriptase, a hepatitis B virus reverse transcriptase, a cauliflower mosaic virus reverse transcriptase, a

bacterial reverse transcriptase, and mutants and variants thereof that are substantially reduced in RNase H activity.

74. (Previously presented) The method according to claim 64, wherein the conditions sufficient to make one or more cDNA molecules comprise use of one or more DNA polymerases, one or more nucleotides and one or more primers.

75. (Previously presented) The method according to claim 74, wherein one or more of the primers are primer-adapters that comprise one or more ligands and one or more cleavage sites.

76. (Previously presented) The method according to claim 64, wherein at least one of the RNA molecules is an mRNA molecule.

77. (Previously presented) The method according to claim 64, wherein at least one of the RNA molecules is polyadenylated.

78. (Previously presented) The method according to claim 64, wherein the one or more RNA molecules is a population of RNA molecules.

79. (Currently amended) A method of making one or more cDNA molecules, comprising:

(a) mixing one or more RNA molecules with (i) one or more polypeptides having reverse transcriptase activity and (ii) at least one primer-adapter nucleic acid molecule wherein the at least one primer-adapter nucleic acid molecule comprises at least one restriction enzyme recognition sequence and at least one biotin moiety, to form a mixture;

- (b) incubating the mixture under conditions sufficient to make one or more double stranded cDNA molecules, wherein one or more of the cDNA molecules comprise at least one primer-adapter nucleic acid molecule;
- (c) contacting one or more of the cDNA molecules with one or more solid supports which comprise avidin and/or streptavidin, to produce one or more solid support-cDNA molecule complexes;
- (d) contacting one or more of the complexes with at least not more than one restriction enzyme that cleaves the at least one restriction enzyme recognition sequence in within the adapter-primer, to produce one or more cleaved cDNA molecules; and
- (e) inserting or ligating one or more of the cleaved cDNA molecules into one or more vectors.

80. (Previously presented) The method according to claim 79, further comprising isolating one or more of the solid support-cDNA molecule complexes.

81. (Previously presented) The method according to claim 79, wherein the solid support is selected from the group consisting of nitrocellulose, diazocellulose, glass, polystyrene, polyvinylchloride, polypropylene, polyethylene, dextran, Sepharose, agar, starch, nylon, beads and microtitre plates.

82. (Previously presented) The method according to claim 79, wherein the solid support is a magnetic bead.

83. (Previously presented) The method according to claim 79, wherein the restriction enzyme is *NotI*.

84. (Previously presented) The method according to claim 79, wherein the one or more polypeptides are selected from the group consisting of a Moloney Leukemia Virus (M-MLV) reverse transcriptase, a Rous Sarcoma Virus (RSV) reverse transcriptase, an Avian Myeloblastosis Virus (AMV) reverse transcriptase, a Rous Associated Virus (RAV) reverse transcriptase, a Myeloblastosis Associated Virus (MAV) reverse transcriptase, a Human Immunodeficiency Virus (HIV) reverse transcriptase, a retroviral reverse transcriptase, a retrotransposon reverse transcriptase, a hepatitis B virus reverse transcriptase, a cauliflower mosaic virus reverse transcriptase, a bacterial reverse transcriptase, and mutants and variants thereof that are substantially reduced in RNase H activity.

85. (Previously presented) The method according to claim 79, wherein the conditions sufficient to make one or more cDNA molecules comprise use of one or more DNA polymerases, one or more nucleotides and one or more primers.

86. (Previously presented) The method according to claim 79, wherein at least one of the RNA molecules is an mRNA molecule.

87. (Previously presented) The method according to claim 79, wherein at least one of the RNA molecules is polyadenylated.

88. (Previously presented) The method according to claim 79, wherein the one or more RNA molecules is a population of RNA molecules.

89. (Currently amended) A method of making one or more cDNA molecules, comprising:

- (a) mixing one or more ~~mRNA RNA~~ molecules with (i) one or more polypeptides having reverse transcriptase activity and (ii) at least one primer-adapter nucleic acid molecule, wherein the at least one primer-adapter nucleic acid molecule comprises one or more ligands and one or more cleavage sites, to form a mixture;
- (b) incubating the mixture under conditions sufficient to make one or more double stranded cDNA molecules, wherein one or more of the cDNA molecules comprise at least one primer-adapter nucleic acid molecule;
- (c) contacting one or more of the cDNA molecules with at least one hapten under conditions sufficient to form one or more hapten-cDNA molecule complexes; **and**
- (d) isolating one or more of the complexes comprising the cDNA molecules; **and**
- (e) cleaving one or more of the isolated complexes with not more than one enzyme that cleaves one or more of the complexes at one or more cleavage sites within the primer-adapter.

90. (Cancelled).

91. (Currently amended) The method according to claim 90 89, further comprising inserting or ligating one or more of the cleaved cDNA molecules into one or more vectors.

92. (Previously presented) The method according to claim 89, wherein the hapten is bound to a solid support.

93. (Previously presented) The method according to claim 92, wherein the solid support is selected from the group consisting of nitrocellulose, diazocellulose, glass, polystyrene, polyvinylchloride, polypropylene, polyethylene, dextran, Sepharose, agar, starch, nylon, beads and microtitre plates.

94. (Previously presented) The method according to claim 92, wherein the solid support is a magnetic bead.

95. (Currently amended) The method according to claim 94 89, wherein the enzyme is *NotI*.

96. (Previously presented) The method according to claim 91, wherein one or more of the cleaved cDNA molecules comprise one sticky end and one blunt end.

97. (Previously presented) The method according to claim 96, wherein the sticky end is a *NotI* sticky end and the vector has a *NotI* compatible end and a blunt end.

98. (Previously presented) The method according to claim 89, wherein the one or more polypeptides are selected from the group consisting of a Moloney Leukemia Virus (M-MLV) reverse transcriptase, a Rous Sarcoma Virus (RSV) reverse transcriptase, an Avian Myeloblastosis Virus (AMV) reverse transcriptase, a Rous Associated Virus (RAV) reverse transcriptase, a Myeloblastosis Associated Virus (MAV) reverse transcriptase, a Human Immunodeficiency Virus (HIV) reverse transcriptase, a retroviral reverse transcriptase, a retrotransposon reverse transcriptase, a hepatitis B virus reverse transcriptase, a cauliflower mosaic virus reverse transcriptase, a

bacterial reverse transcriptase, and mutants and variants thereof that are substantially reduced in RNase H activity.

99. (Previously presented) The method according to claim 89, wherein the conditions sufficient to make one or more cDNA molecules comprise use of one or more DNA polymerases, one or more nucleotides and one or more primers.

100. (Currently amended) The method according to claim 99, wherein one or more of the primers are primer-adapters that comprise one or more ligands and one or more cleavage sites.

101. (Previously presented) The method according to claim 89, wherein at least one of the RNA molecules is an mRNA molecule.

102. (Previously presented) The method according to claim 89, wherein at least one of the RNA molecules is polyadenylated.

103. (Previously presented) The method according to claim 89, wherein the one or more RNA molecules is a population of RNA molecules.

104. (Currently amended) A method of making one or more cDNA molecules, comprising:

(a) mixing one or more mRNA RNA molecules with (i) one or more polypeptides having reverse transcriptase activity and (ii) at least one primer-adapter nucleic acid molecule, wherein the at least one primer-adapter nucleic acid molecule comprises one or more ligands and one or more cleavage sites, to form a mixture;

- (b) incubating the mixture under conditions sufficient to make one or more double stranded cDNA molecules, wherein one or more of the cDNA molecules comprise at least one primer-adapter nucleic acid molecule;
- (c) contacting one or more of the cDNA molecules with at least one hapten under conditions sufficient to form one or more hapten-cDNA molecule complexes; and
- (d) cleaving one or more of the complexes with at least not more than one enzyme that cleaves one or more of the complexes at one or more cleavage sites in within the primer-adapter, , to produce one or more cleaved cDNA molecules.

105. (Previously presented) The method according to claim 104, further comprising isolating one or more of the complexes using a solid support.

106. (Previously presented) The method according to claim 104, further comprising ligating or inserting the cleaved cDNA molecules into one or more plasmids.

107. (Previously presented) The method according to claim 104, wherein the hapten is bound to a solid support.

108. (Currently amended) The method according to claim ~~108~~ 107, wherein the solid support is selected from the group consisting of nitrocellulose, diazocellulose, glass, polystyrene, polyvinylchloride, polypropylene, polyethylene, dextran, Sepharose, agar, starch, nylon, beads and microtitre plates.

109. (Currently amended) The method according to claim ~~108~~ 107, wherein the solid support is a magnetic bead.

110. (Currently amended) The method according to claim 104, wherein the ~~restriction~~ enzyme is *NotI*.

111. (Previously presented) The method according to claim 106, wherein the cleaved cDNA molecules comprise one sticky end and one blunt end.

112. (Currently amended) The method according to claim 111, wherein the sticky end is a *NotI* sticky end and ~~the vector at least one of the plasmids~~ has a *NotI* compatible end and a blunt end.

113. (Previously presented) The method according to claim 104, wherein the one or more polypeptides are selected from the group consisting of a Moloney Leukemia Virus (M-MLV) reverse transcriptase, a Rous Sarcoma Virus (RSV) reverse transcriptase, an Avian Myeloblastosis Virus (AMV) reverse transcriptase, a Rous Associated Virus (RAV) reverse transcriptase, a Myeloblastosis Associated Virus (MAV) reverse transcriptase, a Human Immunodeficiency Virus (HIV) reverse transcriptase, a retroviral reverse transcriptase, a retrotransposon reverse transcriptase, a hepatitis B virus reverse transcriptase, a cauliflower mosaic virus reverse transcriptase, a bacterial reverse transcriptase, and mutants and variants thereof that are substantially reduced in RNase H activity.

114. (Previously presented) The method according to claim 104, wherein the conditions sufficient to make one or more cDNA molecules comprise use of one or more DNA polymerases, one or more nucleotides and one or more primers.

115. (Currently amended) The method according to claim 114, wherein one or more of the primers are primer-adapters that comprise one or more ligands and one or more cleavage sites.

116. (Previously presented) The method according to claim 104, wherein at least one of the RNA molecules is an mRNA molecule.

117. (Previously presented) The method according to claim 104, wherein at least one of the RNA molecules is polyadenylated.

118. (Previously presented) The method according to claim 104, wherein the one or more RNA molecules is a population of RNA molecules.

119. (Currently amended) A method of making one or more cDNA molecules, comprising:

(a) mixing one or more RNA molecules with (i) one or more reverse transcriptases and (ii) one or more primer-adapter nucleic acid molecules, wherein one or more of the primer-adapter nucleic acid molecules comprise at least one restriction enzyme recognition sequence and at least one ligand, to form a mixture;

(b) incubating the mixture under conditions sufficient to make one or more double stranded cDNA molecules, wherein one or more of the cDNA molecules comprise at least one primer-adapter nucleic acid molecule;

(c) contacting one or more of the cDNA molecules with one or more solid supports which comprise one or more haptens, to produce one or more solid support-cDNA molecule complexes;

- (d) cleaving one or more of the complexes with ~~at least~~ not more than one restriction enzyme that cleaves the restriction enzyme recognition sequence ~~in~~ within the primer-adapters, to produce one or more cleaved cDNA molecules; and
- (e) inserting or ligating one or more of the cleaved cDNA molecules into one or more vectors.

120. (Currently amended) The method of claim ~~56~~ 119, wherein the restriction enzyme is *NotI*.

121. (Currently amended) The method of claim ~~56~~ 119, wherein the ligand is a biotin moiety.

122. (Currently amended) The method of claim ~~56~~ 119, wherein the hapten is avidin or streptavidin.

123. (New) The method of claim 64, wherein the one or more polypeptides are selected from the group consisting of a Moloney Leukemia Virus (M-MLV) reverse transcriptase, a Rous Sarcoma Virus (RSV) reverse transcriptase and an Avian Myeloblastosis Virus (AMV) reverse transcriptase.

124. (New) The method of claim 64, wherein the polypeptide is a SuperScript™ reverse transcriptase.

125. (New) The method of claim 64, wherein the ligand is biotin and the hapten is avidin or streptavidin.

126. (New) The method of claim 79, wherein the one or more polypeptides are selected from the group consisting of a Moloney Leukemia Virus (M-MLV) reverse transcriptase, a Rous Sarcoma Virus (RSV) reverse transcriptase and an Avian Myeloblastosis Virus (AMV) reverse transcriptase.

127. (New) The method of claim 79, wherein the polypeptide is a SuperScript™ reverse transcriptase.

128. (New) The method of claim 89, wherein the one or more polypeptides are selected from the group consisting of a Moloney Leukemia Virus (M-MLV) reverse transcriptase, a Rous Sarcoma Virus (RSV) reverse transcriptase and an Avian Myeloblastosis Virus (AMV) reverse transcriptase.

129. (New) The method of claim 89, wherein the polypeptide is a SuperScript™ reverse transcriptase.

130. (New) The method of claim 89, wherein the ligand is biotin and the hapten is avidin or streptavidin.

131. (New) The method of claim 104, wherein the one or more polypeptides are selected from the group consisting of a Moloney Leukemia Virus (M-MLV) reverse transcriptase, a Rous Sarcoma Virus (RSV) reverse transcriptase and an Avian Myeloblastosis Virus (AMV) reverse transcriptase.

132. (New) The method of claim 104, wherein the polypeptide is a SuperScript™ reverse transcriptase.

133. (New) The method of claim 104, wherein the ligand is biotin and the hapten is avidin or streptavidin.

134. (New) The method of claim 119, wherein the one or more reverse transcriptases are selected from the group consisting of a Moloney Leukemia Virus (M-MLV) reverse transcriptase, a Rous Sarcoma Virus (RSV) reverse transcriptase and an Avian Myeloblastosis Virus (AMV) reverse transcriptase.

135. (New) The method of claim 119, wherein the reverse transcriptase is a SuperScript™ reverse transcriptase.